

This Page Is Inserted by IFW Operations
and is not a part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images may include (but are not limited to):

- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

IMAGES ARE BEST AVAILABLE COPY.

**As rescanning documents *will not* correct images,
please do not report the images to the
Image Problems Mailbox.**

THIS PAGE BLANK (USPTO)

PCT

WORLD INTELLECTUAL PROPERTY ORGANIZATION
International Bureau



A61L

27/28

10/24/00
09/695562
Jc917 U.S. PRO

INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁶ : A61L 27/00		(11) International Publication Number: WO 99/58167
A1		(43) International Publication Date: 18 November 1999 (18.11.99)
(21) International Application Number: PCT/US99/08456		(74) Agent: BRUCKNER, John, J.; Wilson Sonsini Goodrich & Rosati, 650 Page Mill Road, Palo Alto, CA 94304-1050 (US).
(22) International Filing Date: 16 April 1999 (16.04.99)		(81) Designated States: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, ARIPO patent (GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).
(30) Priority Data: 09/079,502 14 May 1998 (14.05.98) US		
(63) Related by Continuation (CON) or Continuation-in-Part (CIP) to Earlier Application US 09/079,502 (CIP) Filed on 14 May 1998 (14.05.98)		
(71) Applicant (for all designated States except US): HAYES MEDICAL, INC. [US/US]; Suite 100, 1115 Windfield Way, El Dorado Hills, CA 95762-9623 (US).		
(72) Inventors; and (75) Inventors/Applicants (for US only): DESPRES, Alfred, S., III [US/US]; 2313 Muir Woods Place, Davis, CA 95616 (US). ELWOOD, Eugene, J. [US/US]; 100 Grand Avenue, Cedar Knolls, NJ 07927 (US). AHARONOV, Robert, R. [US/US]; 5 Cayuga Avenue, Rockaway, NJ 07866 (US). EHLERS, Peter [DE/DE]; Zum Walzenberg 15, D-78476 Allensbach (DE). ANDERSEN, Knut [DE/DE]; Schlossberg-Strasse 32A, D-78476 Allensbach-Hegne (DE).		

Published
With international search report.

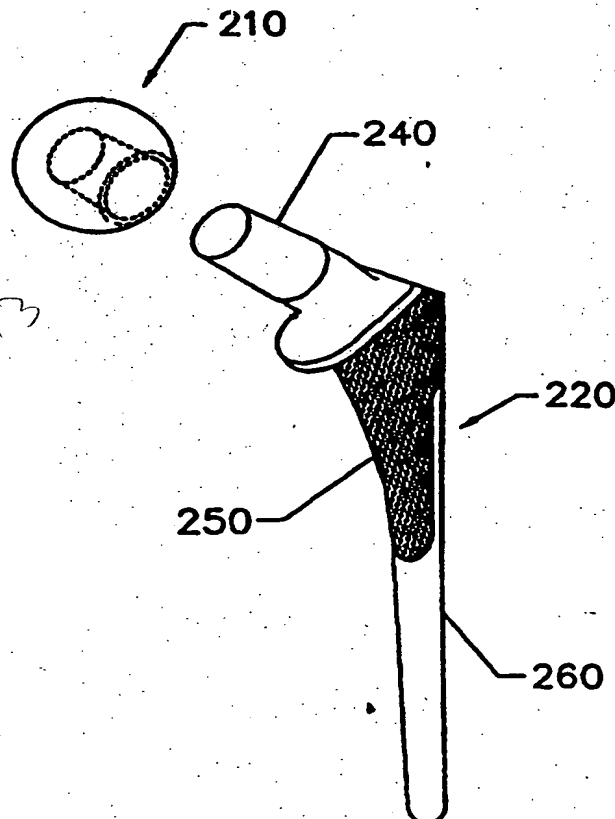
W099/58167

(54) Title: **IMPLANT WITH COMPOSITE COATING**

(57) Abstract

Systems and methods are described for implants with composite coatings to promote tissue in-growth and/or on-growth. An implant includes: a substrate; a structured surface formed on at least a portion of the substrate; and a biocompatible coating deposited on at least a fraction of the structured surface. The systems and methods provide advantages in that the implant has good biocompatibility while the biocompatible coating has good strength.

p7, 11-12, 13 dots



E

FOR THE PURPOSES OF INFORMATION ONLY

Codes used to identify States party to the PCT on the front pages of pamphlets publishing international applications under the PCT.

AL	Albania	ES	Spain	LS	Lesotho	SI	Slovenia
AM	Armenia	FI	Finland	LT	Lithuania	SK	Slovakia
AT	Austria	FR	France	LU	Luxembourg	SN	Senegal
AU	Australia	GA	Gabon	LV	Latvia	SZ	Swaziland
AZ	Azerbaijan	GB	United Kingdom	MC	Monaco	TD	Chad
BA	Bosnia and Herzegovina	GE	Georgia	MD	Republic of Moldova	TG	Togo
BB	Barbados	GH	Ghana	MG	Madagascar	TJ	Tajikistan
BE	Belgium	GN	Guinea	MK	The former Yugoslav Republic of Macedonia	TM	Turkmenistan
BF	Burkina Faso	GR	Greece			TR	Turkey
BG	Bulgaria	HU	Hungary	ML	Mali	TT	Trinidad and Tobago
BJ	Benin	IE	Ireland	MN	Mongolia	UA	Ukraine
BR	Brazil	IL	Israel	MR	Mauritania	UG	Uganda
BY	Belarus	IS	Iceland	MW	Malawi	US	United States of America
CA	Canada	IT	Italy	MX	Mexico	UZ	Uzbekistan
CF	Central African Republic	JP	Japan	NE	Niger	VN	Viet Nam
CG	Congo	KE	Kenya	NL	Netherlands	YU	Yugoslavia
CH	Switzerland	KG	Kyrgyzstan	NO	Norway	ZW	Zimbabwe
CI	Côte d'Ivoire	KP	Democratic People's Republic of Korea	NZ	New Zealand		
CM	Cameroon			PL	Poland		
CN	China	KR	Republic of Korea	PT	Portugal		
CU	Cuba	KZ	Kazakhstan	RO	Romania		
CZ	Czech Republic	LC	Saint Lucia	RU	Russian Federation		
DE	Germany	LJ	Liechtenstein	SD	Sudan		
DK	Denmark	LK	Sri Lanka	SE	Sweden		
EE	Estonia	LR	Liberia	SG	Singapore		

IMPLANT WITH COMPOSITE COATING

BACKGROUND OF THE INVENTION

1. Field of the Invention

The invention relates generally to the field of *in vivo* implants. More particularly, preferred embodiments of the invention are directed to an orthopedic prosthesis having a composite coating to promote tissue in-growth and/or tissue on-growth.

2. Discussion of the Related Art

Prior art implants are known to those skilled in the art. For example, conventional implants are typically composed of stainless steel, cobalt-chrome, or titanium alloy.

A problem with this technology has been that the best substrate materials are not the best materials to be in contact with living tissue. Therefore, what is required is a solution that can provide a biocompatible coating on a substrate.

One approach in an attempt to solve the above-discussed problems involves simply coating an implant substrate with a biocompatible material. However, a disadvantage of this approach is that biocompatible materials are often soft or brittle.

Another disadvantage of this approach has been relatively high cost and/or technical complexity. Therefore, what is also needed is a solution that meets the above-discussed requirements in a more simple and cost effective manner.

Heretofore, the requirements of good substrate properties, and good coating properties referred to above have not been fully met in combination. What is needed is a solution that simultaneously addresses both of these requirements.

SUMMARY OF THE INVENTION

A primary object of the invention is to provide implants with a composite coating to promote tissue in-growth and/or tissue on-growth. Another primary object of the invention is to provide a composition that can be used as the composite coating. Another primary object of the invention is to provide implants having a composite coating that is located only on surface areas of the implant that are to be fixed with tissue in-growth and/or on-growth for

stability. Another primary object of the invention is to provide methods of making the orthopedic implant.

In accordance with these objects, there is a particular need for an implant with a composite coating. Thus, it is rendered possible to simultaneously satisfy the above-discussed requirements of good substrate properties and good biocompatible coating properties, which, in the case of the prior art, are mutually contradicting and cannot be easily satisfied.

A first aspect of the invention is implemented in an embodiment that is based on an implant, comprising: a substrate; a structured surface defined by at least a portion of said substrate; and a biocompatible coating deposited on at least a fraction of said structured surface. A second aspect of the invention is implemented in an embodiment that is based on a composition for an implant, comprising: a biocompatible material coated on a structured surface defined by a substrate. A third aspect of the invention is implemented in an embodiment that is based on an implant, comprising: a substrate; a structured surface defined by a portion of said substrate; and a biocompatible coating deposited on at least a fraction of said structured surface, wherein said portion of said substrate is to be fixed with tissue in-growth and/or on-growth for stability. A fourth aspect of the invention is implemented in an embodiment that is based on a method of forming a composite coating, comprising: depositing a biocompatible coating on a structured surface defined by at least a portion of a surface area of a substrate.

These, and other, objects and aspects of the invention will be better appreciated and understood when considered in conjunction with the following description and the accompanying drawings. It should be understood, however, that the following description, while indicating preferred embodiments of the invention and numerous specific details thereof, is given by way of illustration and not of limitation. Many changes and modifications may be made within the scope of the invention without departing from the spirit thereof, and the invention includes all such modifications.

BRIEF DESCRIPTION OF THE DRAWINGS

A clear conception of the advantages and features constituting the invention, and of the components and operation of model systems provided with the invention, will become

more readily apparent by referring to the exemplary, and therefore nonlimiting, embodiments illustrated in the drawings accompanying and forming a part of this specification, wherein like reference characters (if they occur in more than one view) designate the same parts. It should be noted that the features illustrated in the drawings are not necessarily drawn to scale.

FIG. 1 illustrates a schematic perspective view of an orthopedic implant, representing an embodiment of the invention.

FIG. 2 illustrates a schematic perspective view of an orthopedic implant, representing an embodiment of the invention.

FIGS. 3a-3b illustrate micrograph views of a physical vapor deposited coating of titanium on a structured surface of cobalt-chrome alloy, representing an embodiment of the invention.

FIG. 4 illustrates a schematic view of a coating on a porous structured surface, representing an embodiment of the invention.

FIG. 5 illustrates a schematic view of an apparatus and process for coating a structured surface, representing an embodiment of the invention.

FIGS. 6a-6b illustrate scanning electron micrographs of a biocompatible coating on CoCrMo particles after an electrochemical test, representing an embodiment of the invention.

FIGS. 7a-7d illustrate scanning electron micrographs of CoCrMo particles without the biocompatible coating after the electrochemical test, representing an embodiment of the invention.

FIGS. 8a-8b illustrate SEM of CoCrMo particles without coating before test, representing an embodiment of the invention.

DESCRIPTION OF PREFERRED EMBODIMENTS

The invention and the various features and advantageous details thereof are explained more fully with reference to the nonlimiting embodiments that are illustrated in the accompanying drawings and detailed in the following description. Descriptions of well known components and processing techniques are omitted so as not to unnecessarily obscure the invention in detail.

The context of the invention is providing an implant to be positioned *in vivo* during surgery, especially orthopedic surgery to replace a joint, such as, for example, a knee joint or a hip joint. Thus, the implant can be used in a method for orthopedic surgery that includes surgically positioning the implant within a vertebra in need thereof. If bone growth is facilitated, the implant can be termed part of an osteoconductive process that includes contacting a bone under *in vivo* conditions with the implant.

Referring to the drawings, a detailed description of preferred embodiments of the invention is provided with respect to FIGS. 1 through 5. The invention is not limited to the specific embodiments illustrated in FIGS. 1-5.

Referring now to FIG. 1, a component 110 of an artificial knee joint is depicted. The component 110 includes a bearing surface 120 and a tissue fixation surface 130. The bearing surface 120 is for sliding. The tissue fixation surface 130 is for tissue in-growth and/or tissue on-growth. The tissue fixation surface 130 can be a coating that is deposited on at least a fraction of a structured surface that is defined by, or formed on, or composed by at least a portion of a substrate.

The substrate can be composed, or formed, of, for example, carbon-composite, stainless steel, cobalt-chromium, titanium alloy, tantalum, and/or ceramic, and combinations thereof. The structured surface can be defined by, or composed of, or formed of a material that includes a plurality of particles that are sintered together to form a continuous porous phase. Alternatively, the structured surface can be prepared by at least one method selected from the group consisting of flame spraying, acid etching, grit blasting, casting-in, forging-in, laser texturing, and micromachining.

The coating should be a biocompatible coating. The biocompatible coating can include one, or more, of titanium, tantalum, carbon, calcium phosphate, zirconium, niobium, hafnium, hydroxyapatite, tissue in-growth and/or on-growth facilitating proteins, and combinations thereof. If carbon is used, it can optionally be diamond-like carbon, pyrolytic carbon, amorphous diamond-like carbon, and combinations thereof.

It can be advantageous that the coating be more biocompatible than the structured surface. Similarly, it can be advantageous that the coating be more biocompatible than the substrate. One aspect, albeit optional, of biocompatibility is softness. It can be advantageous that the coating be softer than the structured surface. Similarly, it can be advantageous that the coating be softer than the substrate. Although the preferred

embodiment shown in FIG. 1 includes a coating located on specific portions of the substrate, it is within the level of ordinary skill in the art after having knowledge of the invention disclosed herein to coat any surface(s) of the substrate.

Referring now to FIG. 2, an artificial hip joint is depicted. The hip joint includes a femoral head 210 and distal portion 220. The femoral head 210 includes a joint motion surface 230 for bearing and sliding. The distal portion 220 includes a taper connection 240, a tissue fixation surface 250, and a distal stem 260. The tissue fixation surface 250 can be a coating that is deposited on at least a fraction of a structured surface that is defined by, or formed on, or composed by at least a portion of a substrate. As noted above, the coating should be a biocompatible coating. The biocompatible coating can include multi-layers. These multi-layers can be nano-layers. Although the preferred embodiment shown in FIG. 2 includes the coating located on a specific portion of the substrate, it is within the level of ordinary skill in the art after having knowledge of the invention disclosed herein to coat any surface(s) of the substrate.

Referring now to FIGS. 3a-3b, two images of a structured surface coated according to the invention are depicted. FIGS. 3a-3b are two images of the same sample acquired at different locations. FIG. 3a illustrates a coating 310 on a structured surface 320 at an outermost location. The structure surface 320 is defined by a sintered layer having interconnected porosity. The thickness of the coating 310 is approximately 10.78 microns at this outermost location of the structured surface 320. FIG. 3b illustrates the coating 310 on the structured surface 320 at an innermost location. The thickness of the coating is approximately 2.0 microns at this innermost location of the structured surface 320.

Still referring to FIGS. 3a-3b, a coating 310 is adhered to a structured surface 320. In this particular embodiment, the coating 310 is composed of a first material that includes titanium. The coating is preferably a biocompatible coating. In this particular embodiment, the structured surface 320 is defined by a second material that includes cobalt and chrome. In this particular embodiment, the structured surface was conditioned by cathodic arc ion plating of titanium before the coating 310 was deposited using the same apparatus used to effect the ion plating. In this particular embodiment, the average thickness of the coating 310 is approximately 10.78 microns. If the structured surface 320 includes crevices and/or undercuts 330, the biocompatible material that composes the coating 310 can coat the crevices and/or undercuts 330 in the structured surface.

It can be advantageous if the biocompatible material conforms to the crevices and/or undercuts and thereby defines a textured (e.g., rough) topology at the upper surface of the biocompatible material. This is advantageous because such a topology gives tissue a better hold via tissue in-growth and/or on-growth. Further, if there are pores in and/or beneath the structured surface the, biocompatible material can coat the pores. It can be advantageous if the biocompatible material coats interconnected pores located beneath the structured surface and thereby define voids (pores) and/or interconnected pores in which tissue in-growth and/or on-growth can occur.

The effect of cathodic arc ion plating with a high bias voltage is to cause an intermixing of titanium into the cobalt-chrome substrate due to high ion energies. It can be appreciated that there is an inter-penetration of the coating material into the structured surface material between the interface and the bulk of the material that composes the structured surface. Specifically, the amount of coating material that composes the substrate that defines the structured surface decreases as the depth from the interface increases. When the bias voltage is increased, the inter-penetration depth increases. The depth of this intermixing can range from approximately 0.5 nm to approximately 500 nm, preferably approximately 50 nm to approximately 100 nm. When the bias voltage is sufficient, substantial inter-penetration occurs. This result is advantageous because the inter-penetration improves the adhesion of the coating to the structured surface, thereby minimizing flaking, peeling, and other disruptions of the coating. This is very important for at least the following two reasons. First, the coating will be subject to tissue in-growth and/or on-growth and, therefore, can be subject to dynamic loading from adjacent tissue structures, such as, for example, bones. By improving the adhesion, the coating is better able to withstand loading. Second, the implant may be expected to remain *in vivo* for many years and it is highly desirable that all of the implant remain fixed in place. By improving the adhesion of the coating to the structured surface, the long term stability of the coating is enhanced.

Referring now to FIG. 4, a porous structured surface 410 with a coating 420 is depicted. The coating 420 includes titanium and can be deposited via physical vapor deposition with a gas, preferably an inert gas, such as, for example, argon and/or helium. It can be advantageous to use argon as the gas because it is inert and has a relatively high atomic weight. Significantly, the coating 420 covers portions of a structured surface defined

by an interconnected pore 430 that is hidden from line-of-sight deposition (i.e., the pore 430 cannot be seen from the point of view of the deposition source). The structured surface is also defined by a plurality of particles covering a substrate 440 (i.e., the structured surface is defined by the interface between the substrate 440 and the plurality of particles).

- 5 Nevertheless, the surface defined by the interconnected pore 430 is coated and provides an area for tissue in-growth and/or on-growth because an inert gas is present in the deposition chamber while the coating 420 is being applied. Although the preferred embodiment shown in FIG. 4 includes the coating of the surface defined by the interconnected pore 430, it is within the level of ordinary skill in the art after having knowledge of the invention disclosed
10 herein to coat any undercut, or vertical, or line-of-sight hidden surface area.

FIG. 4 demonstrates substantially improved results that are unexpected. Specifically, the coating of internal pores demonstrates the significant unexpected advantageous result that when an inert gas is present during the deposition process, the coating is deposited on line-of-sight hidden surfaces (e.g. interconnected pores). It can be appreciated that the
15 structured surface 410 is coated because the exterior of the particles appears rough. Normally, in the case of an uncoated particle, the perimeter would be smooth and more nearly circular. The roughness is the coating. This result is advantageous because it significantly improves adhesion of the surrounding tissue to the implant. Adhesion is significantly improved because tissue on-growth and/or in-growth can take place on
20 undercuts, crevices, cul de sacs, conduits, caves, tunnels, and interconnected pores that are hidden from line-of-sight deposition, thereby significantly improving the strength of the connection between the surrounding tissue and the implant. The strength of the connection is significantly improved because the tissue grows into the undercuts, crevices, cul de sacs, conduits, caves, tunnels, and interconnected pores creating a tissue structure that interlocks
25 with the structured surface on a macroscopic level. It can be advantageous if the coating covers a continuous length of a void structure (e.g., undercuts, crevices, cul de sacs, conduits, caves, tunnels, interconnected pores, etc.) that is open to adjacent tissue in at least two places. For example, if the tissue grows through a tunnel, the strength of the connection will be based not only on the interface adhesion between the wall of the tunnel and the tissue,
30 but also on the inherent mechanical strength of the loop of tissue that is routed through the tunnel.

The particular manufacturing process used for depositing the coating should be inexpensive and reproducible. Conveniently, the deposition of the coating can be carried out by using any vapor deposition method. Vapor deposition methods include chemical vapor deposition (CVD) (e.g., plasma assisted CVD) and physical vapor deposition (PVD) (e.g., arc evaporation, e-beam, molten pool, sputtering, evaporative ion plating, and cathodic arc ion plating). It is preferred that the process be a physical vapor deposition. For the manufacturing operation, it is moreover an advantage to employ an arc evaporation physical vapor deposition method.

FIG. 5 depicts an arc evaporation physical vapor deposition apparatus and method for coating a structured surface on a substrate. A water cooled chamber 510 functions as an anode. A vacuum pump 520 and a conduit 530 for a neutral gas and a reactive gas are connected to the chamber 510. The chamber includes a plurality of evaporators 540 that function as cathodes. Each of the evaporators includes a source of a material 545 from which the coating is to be formed (e.g., titanium). An arc power supply 550 can be connected to each of the plurality of evaporators 540 (only a single connection is shown in FIG. 5). Each of the evaporators 540 can generate a plasma 560 that includes a high number of ions together with electrons and neutral vapor atoms. (It should be noted that the plasma 560 is represented schematically for clarity.) The plasma 560 impinges upon a substrate 570 that is connected to a bias (-) power supply. By increasing the bias, the ions are accelerated toward the substrate more rapidly. The apparatus can also include one, or more, structures to steer and/or filter the plasma such as, for example internal and/or external magnets.

Still referring to FIG. 5, the substrate 570 includes a structured surface (not shown) onto which the coating can be deposited. Portions of the substrate that are not to be coated can be masked with a mask material that can be removed after the deposition of the coating is finished. In general, the coating can be formed by any thin film technique. Thin film technique include physical vapor deposition and chemical vapor deposition, and combinations thereof.

Still referring to FIG. 5, the method of using the illustrated apparatus begins by loading the substrate 570 into the chamber 510. A vacuum is then created in the chamber 510 using the vacuum pump 520. A test for leaks is then conducted. A conditioning and heating subprocess can then be performed. One reason to conduct this subprocess is to heat and remove any oxides from the structured surface. This subprocess can be based on radiant

heating. This subprocess can also be based on glow discharge with a high bias voltage to effect an ion bombardment by a gas, or gas mixture. This subprocess can also be based on ion bombardment with the material from which the coating is to be formed (e.g., titanium) with a high bias voltage (e.g., from approximately 1000 to approximately 1500 volts). This latter technique causes an intermixing of the coating material (e.g., titanium) into the material that defines the structured surface (e.g., CoCr). After the conclusion of the subprocess, the coating is applied. The bias voltage is lowered to deposit and build up the coating. By adding a gas, preferably an inert gas, such as, for example, argon or helium (most preferably argon), full coverage of undercuts and full coverage within pore structures can be obtained.

While not being limited to any particular theory, it is believed that the presence of the gas reduces the average velocity and increases the scattering of the coating vapor as it moves from the cathode to the structured surface, thereby allowing better migration of the coating vapor inside the undercuts, crevices, cul de sacs, conduits, caves, tunnels, interconnected pores, etc., before the vapor is deposited (i.e., fixed) on the structured surface. This relationship may be due to an increase in scattered vector components representing the speed and direction of the vapor molecules due to collisions between the vapor molecules and the gas molecules, with a corresponding decrease in the magnitude of a primary vector representing the average speed and velocity of all the vapor molecules.

Still referring to FIG. 5, the process temperature can range from approximately 500 to approximately 900 deg. F. The operating pressure during radiant heating can range from approximately 1.0 millitorr to approximately 0.01 millitorr. The operating pressure during the bombard phase (e.g., titanium without any dampening gas) can range from approximately 10.0 millitorr to approximately 0.001 millitorr. The operating pressure during the coating phase (e.g., titanium with optional dampening gas) can range from approximately less than 1 millitorr to approximately 100 millitorr. The arc voltage can range from approximately 10 to approximately 100 volts/evaporator. The bias voltage can range from approximately 1000 to approximately 1500 volts during bombardment, and from approximately 10 to approximately 500 volts during coating.

However, the particular manufacturing process used for depositing the coating is not essential to the invention as long as it provides the described biocompatible coating. Normally those who make or use the invention will select the manufacturing process based

upon tooling and energy requirements, the expected application requirements of the final product, and the demands of the overall manufacturing process.

Referring now to FIGS. 6a-6b, scanning electron micrographs of a biocompatible coating including titanium on a structured surface defined by CoCrMo particles after an electrochemical test (ASTM F746) are depicted. FIG. 6a was acquired at x300. FIG. 6b was acquired at x700.

Referring now to FIGS. 7a-7d, electron micrographs of a structured surface defined by CoCrMo particles without the biocompatible coating after the electrochemical test (ASTM F746) are depicted. FIG. 7b was acquired at x120. FIG. 7d was acquired at x650.

By comparing the results shown in FIGS. 7a-7b with the results shown in FIGS. 6a-6b, it can be appreciated that the biocompatible coating significantly inhibits corrosion. For example, FIGS. 7a-7d show significant evidence of intergranular corrosion attacks while FIGS. 6a-6b do not show significant evidence of intergranular corrosion attacks.

Referring now to Figs. 8a-8b, electron micrographs of a structured surface defined by CoCrMo particles without the biocompatible coating before the electrochemical test (ASTM F746) (i.e., before any corrosion test) are depicted. FIG. 8a was acquired at x120. FIG. 8b was acquired at x600.

The particular material used for the substrate should be strong. Some examples of substrate materials include ASTM F67, F75, F90, F136, F138, F560, F562, F620, F621, F799, F961, F1058, F1295, F1341, F1472, F1537, and F1586. These are all grades of titanium, CoCr, stainless steel, tantalum, or alloy(s) thereof. It is preferred that the substrate material be a cobalt-chrome alloy. The substrate can be manufactured as a wrought, forged, or cast form.

The particular material used to define the structured surface should also be strong. It is preferred that the material used to define the structured surface be the same as the material used for the substrate (e.g., cobalt-chrome). However, the material used to define the structured surface can be different and include any one or more of, for example, titanium, CoCr, stainless steel, and tantalum, or alloy(s) thereof. The structured surface can be manufactured in various forms, including, but not limited to, sintered, flame sprayed, acid etched, grit blasted, cast-in, forged-in, laser textured, or micromachined form. For the manufacturing operation, it is an advantage to employ a partially sintered (porous) structure

obtained by forming a layer of a mixture that contains a plurality of particles and a binder, and optionally a sacrificial filler.

For non-interconnecting (nonporous) structures, the surface roughness (R_a or R_z) of the structured surface can range from approximately 100 to approximately 40,000 μ /inch, preferably from approximately 3,000 to approximately 30,000 μ /inch. When the structured surface is defined by a porous layer, the layer can have a thickness from approximately 0.010 to 0.080 inch deep, preferably from approximately 0.010 to 0.060 inch deep. In the case of porous structures, the pore sizes can be from approximately 50 to approximately 500 microns, preferably from approximately 200 to 450 microns.

The particular material used for the coating should be biocompatible and provide a suitable surface for tissue in-growth and/or on-growth. It is preferred that the coating be titanium. However, the coating can include one or more of, for example, titanium, tantalum, calcium phosphate, hydroxyapatite, and tissue in-growth and/or on-growth promoting proteins, or alloys thereof.

When the thickness of the coating is excessively thin, parts of the structured surface may not be covered. On the other hand, when the thickness of the coating is excessively high, the strength of the coating may go down due to residual, thermal induced stress. For the specific embodiment of pure titanium on a structured surface defined by a sintered plurality of particles with interconnected porosity, in the deepest region (e.g., bottom of the lowest pore), the minimum coating thickness can be approximately 1 micron. In the outermost region (e.g., top of the highest particle), the maximum coating thickness can be approximately 20 microns.

However, the particular materials selected for coating and substrate are not essential to the invention, as long as they provide the described functions. Normally, those who make or use the invention will select the best commercially available materials based upon the economics of cost and availability, the expected specific application requirements of the final product, and the demands of the overall manufacturing process.

The disclosed embodiments show a continuous coating deposited in one or more zones across at least a portion of a structured surface as the structure for performing the function of facilitating tissue in-growth and/or on-growth. However, the structure for promoting tissue in-growth and/or on-growth can be any other structure capable of

performing the function of achieving tissue adhesion, including, by way of example, a pattern such as an array of dots, or lines, or any other geometric pattern.

While not being limited to any particular performance indicator or diagnostic identifier, preferred embodiments of the invention can be identified one at a time by testing for the presence of high adhesion between the coating and the structured surface. The test for the presence of high adhesion can be carried out without undue experimentation by the use of simple and conventional tribological and mechanical experiments. Among the other ways in which to seek embodiments having the attribute of high adhesion guidance toward the next preferred embodiment can be based on the presence of high coating coverage of undercut surfaces (e.g., hidden interconnected pores). The test for the presence of high coating coverage of undercut surfaces can be carried out without undue experimentation by the use of simple and conventional optical and/or electron microscope imaging techniques.

Practical Applications of the Invention

A practical application of the invention that has value within the technological arts is the fabrication of an implant for the replacement of a knee joint or a hip joint. Further, the invention is useful in conjunction with other implants (such as pins used for the purpose of joining bones), or the like. There are virtually innumerable uses for the invention, all of which need not be detailed here.

Advantages of the Invention

An implant, representing an embodiment of the invention, can be cost effective and advantageous for at least the following reasons. The implants are simple and economical to implement. The implants have increased strength due to the structured surface being defined by a high strength material and better biocompatibility due to the use of materials for the coating that are more biocompatible than the substrate. A single implant can possess different surface areas that are individually optimized with i) some areas bearing tissue in-growth and/or on-growth composite coating, ii) some areas exhibiting the properties of the substrate material(s), and iii) some areas bearing hard surfaces formed by treatments, such as, for example, ion bombardment.

All the disclosed embodiments of the invention described herein can be realized and practiced without undue experimentation. Although the best mode of carrying out the invention contemplated by the inventors is disclosed above, practice of the invention is not

limited thereto. Accordingly, it will be appreciated by those skilled in the art that the invention may be practiced otherwise than as specifically described herein.

For example, the individual components need not be formed in the disclosed shapes, or assembled in the disclosed configuration, but could be provided in virtually any shape, and assembled in virtually any configuration. Further, the individual components need not be fabricated from the disclosed materials, but could be fabricated from virtually any suitable materials. Further, although the implant described herein is a physically separate module, it will be manifest that the implant may be integrated into additional apparatus with which it is associated. Furthermore, all the disclosed elements and features of each disclosed embodiment can be combined with, or substituted for, the disclosed elements and features of every other disclosed embodiment except where such elements or features are mutually exclusive.

It will be manifest that various additions, modifications and rearrangements of the features of the invention may be made without deviating from the spirit and scope of the underlying inventive concept. It is intended that the scope of the invention as defined by the appended claims and their equivalents cover all such additions, modifications, and rearrangements. The appended claims are not to be interpreted as including means-plus-function limitations, unless such a limitation is explicitly recited in a given claim using the phrase "means-for." Expedient embodiments of the invention are differentiated by the appended subclaims.

CLAIMS

What is claimed is:

1. An implant, comprising:
a substrate;
a structured surface formed on at least a portion of said substrate; and
a biocompatible coating deposited on at least a fraction of said structured surface.
2. The implant of claim 1, wherein said biocompatible coating is more biocompatible than said structured surface.
3. The implant of claim 1, wherein said biocompatible coating is more biocompatible than said substrate.
4. The implant of claim 1, wherein said biocompatible coating is softer than said structured surface.
5. The implant of claim 1, wherein said biocompatible coating is softer than said substrate.
6. The implant of claim 1, wherein said coating is formed by a thin film technique.
7. The implant of claim 6, wherein said thin film technique includes at least one deposition process selected from the group consisting of physical vapor deposition and chemical vapor deposition.
8. The implant of claim 1, wherein said structured surface includes a plurality of undercuts and said biocompatible material coats said plurality of undercuts in said structured surface.
9. The implant of claim 8, wherein said structured surface is porous and said biocompatible material coats interconnected pores beneath said structured surface.

10. The implant of claim 1, wherein said substrate includes at least one material selected from the group consisting of carbon-composite, stainless steel, cobalt-chromium, titanium alloy, tantalum, and ceramic.
11. The implant of claim 1, wherein said structured surface is defined by a material that includes a plurality of particles that are sintered together to form a continuous porous phase.
12. The implant of claim 1, wherein said structured surface is prepared by at least one method selected from the group consisting of sintering, flame spraying, acid etching, grit blasting, casting-in, forging-in, laser texturing, and micromachining.
13. The implant of claim 1, wherein said biocompatible coating includes at least one material selected from the group consisting of titanium, tantalum, carbon, calcium phosphate, zirconium, niobium, hafnium, hydroxyapatite, and tissue in-growth and/or on-growth facilitating proteins.
14. The implant of claim 1, wherein said biocompatible coating includes multi-layers.
15. The implant of claim 1, wherein said biocompatible coating includes nano-layers.
16. The implant of claim 1, wherein said implant is an orthopedic prosthesis.
17. A method for orthopedic surgery which comprises surgically positioning said implant of claim 1 within a vertebrate in need thereof.
18. An osteoconductive process, comprising contacting a bone under *in vivo* conditions with said implant of claim 1.
19. A composition for an implant, comprising:
a biocompatible material coated on a structured surface defined by a substrate.

20. The composition of claim 19, wherein said biocompatible coating is more biocompatible than said structured surface.
21. The composition of claim 19, wherein said biocompatible coating is more biocompatible than said substrate.
22. The composition of claim 19, wherein said biocompatible coating is softer than said structured surface.
23. The composition of claim 19, wherein said biocompatible coating is softer than said substrate.
24. The composition of claim 19, wherein said coating is formed by a thin film technique.
25. The composition of claim 24, wherein said thin film technique includes at least one deposition process selected from the group consisting of physical vapor deposition and chemical vapor deposition.
26. The composition of claim 19, wherein said structured surface includes a plurality of undercuts and said biocompatible material coats said plurality of undercuts in said structured surface.
27. The composition of claim 26, wherein said structured surface is porous and said biocompatible material coats interconnected pores beneath said structured surface.
28. The composition of claim 19, wherein said substrate includes at least one material selected from the group consisting of carbon-composite, stainless steel, cobalt-chromium, titanium alloy, tantalum, and ceramic.
29. The composition of claim 19, wherein said structured surface is defined by a material that includes a plurality of particles that are sintered together to form a continuous porous phase.

30. The composition of claim 19, wherein said structured surface is prepared by at least one method selected from the group consisting of sintering, flame spraying, acid etching, grit blasting, casting-in, forging-in, laser texturing, and micromachining.

31. The composition of claim 19, wherein said biocompatible coating includes at least one material selected from the group consisting of titanium, tantalum, carbon, calcium phosphate, zirconium, niobium, hafnium, hydroxyapatite, and tissue in-growth and/or on-growth facilitating proteins.

32. The composition of claim 19, wherein said biocompatible coating includes multi-layers.

33. The composition of claim 19, wherein said biocompatible coating includes nano-layers.

34. An osteoconductive process, comprising contacting a bone under *in vivo* conditions with said composition of claim 19.

35. An orthopedic implant, comprising said composition of claim 19.

36. A method for orthopedic surgery which comprises positioning said composition of claim 19 within a vertebra in need thereof.

37. An implant, comprising:
a substrate;
a structured surface formed on a portion of said substrate; and
a biocompatible coating deposited on at least a fraction of said structured surface,
wherein said portion of said substrate is to be fixed with tissue in-growth and/or on-growth for stability.

38. The implant of claim 37, wherein said biocompatible coating is more biocompatible than said structured surface.

39. The implant of claim 37, wherein said biocompatible coating is more biocompatible than said substrate.
40. The implant of claim 37, wherein said biocompatible coating is softer than said structured surface.
41. The implant of claim 37, wherein said biocompatible coating is softer than said substrate.
42. The implant of claim 37, wherein said coating is formed by a thin film technique.
43. The implant of claim 42, wherein said thin film technique includes at least one deposition process selected from the group consisting of physical vapor deposition and chemical vapor deposition.
44. The implant of claim 37, wherein said structured surface includes a plurality of undercuts and said biocompatible material coats said plurality of undercuts in said structured surface.
45. The implant of claim 44, wherein said structured surface is porous and said biocompatible material coats interconnected pores beneath said structured surface.
46. The implant of claim 37, wherein said substrate includes at least one material selected from the group consisting of carbon-composite, stainless steel, cobalt-chromium, titanium alloy, tantalum, and ceramic.
47. The implant of claim 37, wherein said structured surface is defined by a material that includes a plurality of particles that are sintered together to form a continuous porous phase.
48. The implant of claim 37, wherein said structured surface is prepared by at least one method selected from the group consisting of sintering, flame spraying, acid etching, grit blasting, casting-in, forging-in, laser texturing, and micromachining.

49. The implant of claim 37, wherein said biocompatible coating includes at least one material selected from the group consisting of titanium, tantalum, carbon, calcium phosphate, zirconium, niobium, hafnium, hydroxyapatite, and tissue in-growth and/or on-growth facilitating proteins.
50. The implant of claim 37, wherein said biocompatible coating includes multi-layers.
51. The implant of claim 37, wherein said biocompatible coating includes nano-layers.
52. The implant of claim 37, wherein said implant is an orthopedic prosthesis.
53. A method for orthopedic surgery which comprises surgically positioning said implant of claim 37 within a vertebrate in need thereof.
54. An osteoconductive process, comprising contacting a bone under *in vivo* conditions with said implant of claim 37.
55. A method of forming a composite coating, comprising:
depositing a biocompatible coating on a structured surface that composes at least a portion of a surface area of a substrate.
56. The method of forming a composite coating according to claim 55, further comprising:
covering said portion of said surface area of said substrate with a mixture including a plurality of particles, said plurality of particles including a first material; and
sintering said plurality of particles to produce a porous structure,
wherein said biocompatible coating includes a second material that is different from said first material.
57. The method of claim 55, wherein depositing said biocompatible coating includes depositing a coating that is more biocompatible than said structured surface.

58. The method of claim 55, wherein depositing said biocompatible coating includes depositing a coating that is more biocompatible than said substrate.
59. The method of claim 55, wherein depositing said biocompatible coating includes depositing a coating that is softer than said structured surface.
60. The method of claim 55, wherein depositing said biocompatible coating includes depositing a coating that is softer than said substrate.
61. The method of claim 55, wherein said depositing said biocompatible coating includes forming said biocompatible coating with a thin film technique.
62. The method of claim 61, wherein said thin film technique includes at least one deposition process selected from the group consisting of physical vapor deposition and chemical vapor deposition.
63. The method of claim 55, further comprising, before the step of depositing, forming a plurality of undercuts in said structured surface, and, wherein depositing said biocompatible coating includes coating said plurality of undercuts in said structured surface.
64. The method of claim 63, wherein forming a plurality of undercuts includes forming interconnected pores beneath said structured surface, and, wherein depositing said biocompatible coating includes coating said interconnected pores.
65. The method of claim 55, further comprising, before the step of depositing, providing said substrate from at least one material selected from the group consisting of carbon-composite, stainless steel, cobalt-chromium, titanium alloy, tantalum, and ceramic.
66. The method of claim 55, further comprising, before the step of depositing, providing said structured surface from a material that includes a plurality of particles that are sintered together to form a continuous porous phase.

67. The method of claim 55, further comprising, before the step of depositing, preparing said structured surface by at least one method selected from the group consisting of sintering, flame spraying, acid etching, grit blasting, casting-in, forging-in, laser texturing, and micromachining.
68. The method of claim 55, wherein depositing said biocompatible coating includes depositing at least one material selected from the group consisting of titanium, tantalum, carbon, calcium phosphate, zirconium, niobium, hafnium, hydroxyapatite, and tissue in-growth and/or on-growth facilitating proteins.
69. The method of claim 55, wherein depositing said biocompatible coating include depositing multi-layers.
70. The method of claim 69, wherein depositing said biocompatible coating includes depositing nano-layers.
71. A method for orthopedic surgery which comprises surgically positioning an implant that includes the composite coating made by the method of claim 55 within a vertebrae in need thereof.
72. An osteoconductive process, comprising contacting a bone under *in vivo* conditions with an implant including a composite coating made by the method of claim 55.
73. An implant including a composite coating made by the method of claim 55.
74. An orthopedic prosthesis including a composite coating made by the method of claim 55.
75. The method of claim 55, further comprising, after the step of depositing, treating another portion of said surface area with ion bombardment.

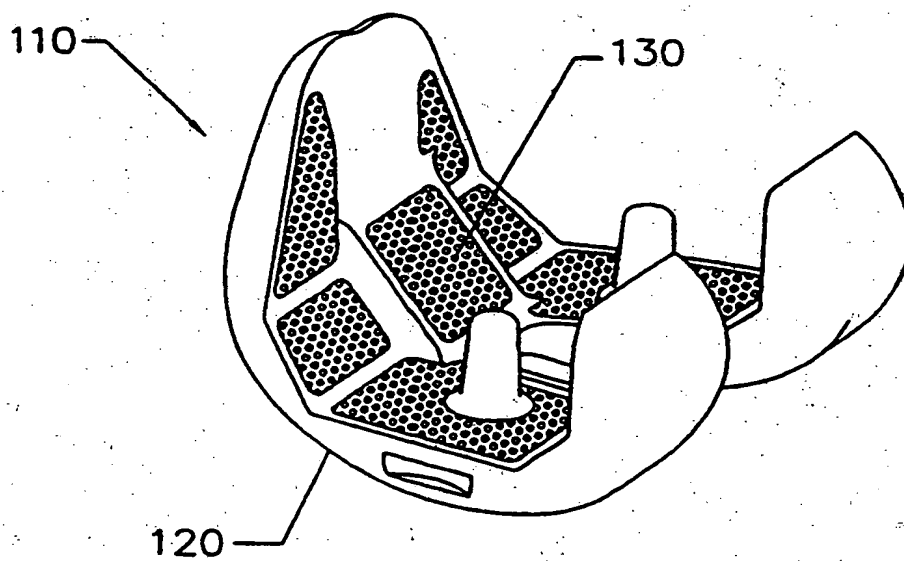


Fig. 1

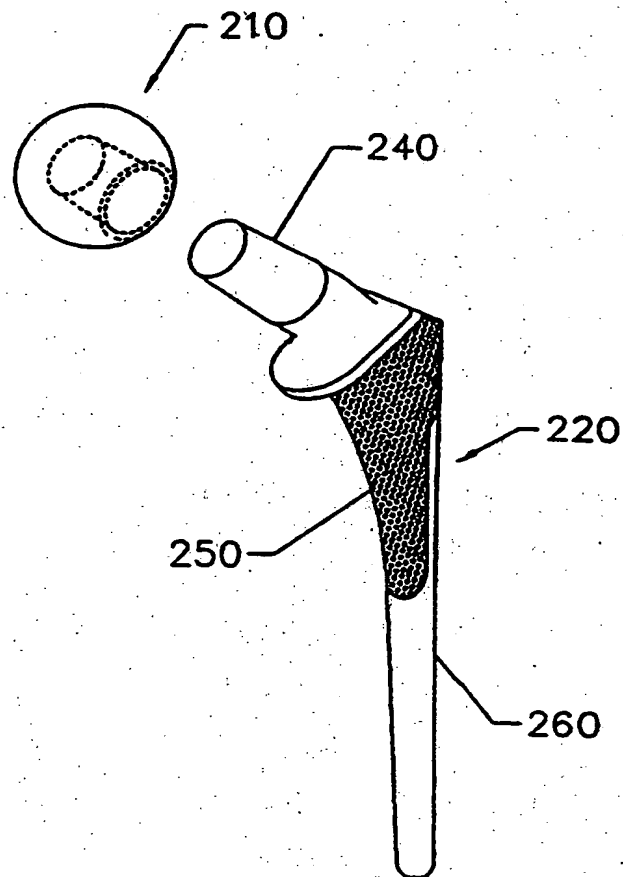


Fig. 2

Fig 3A.

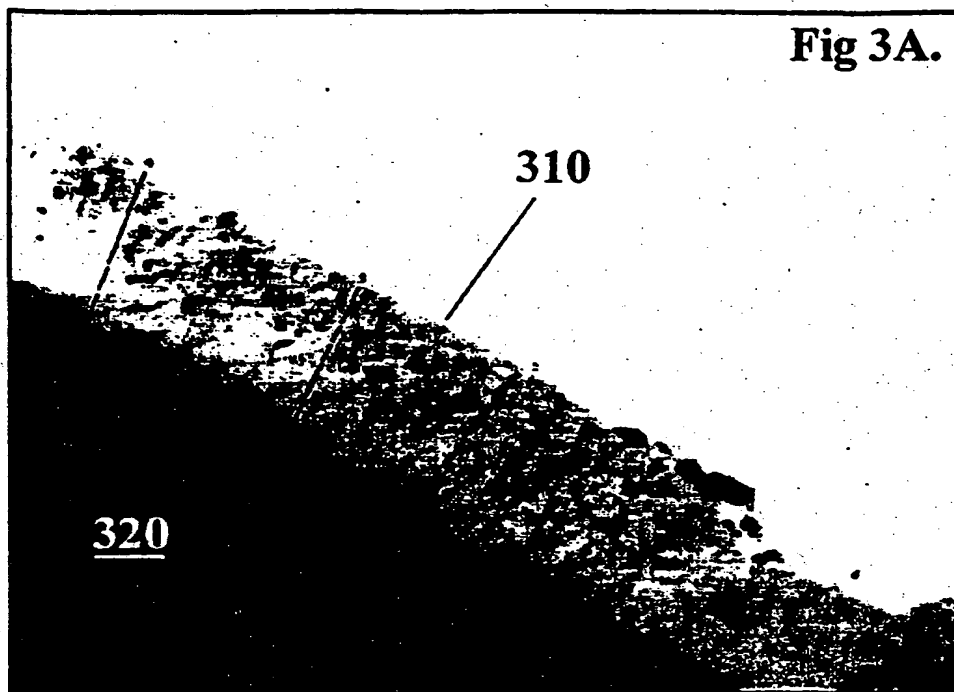
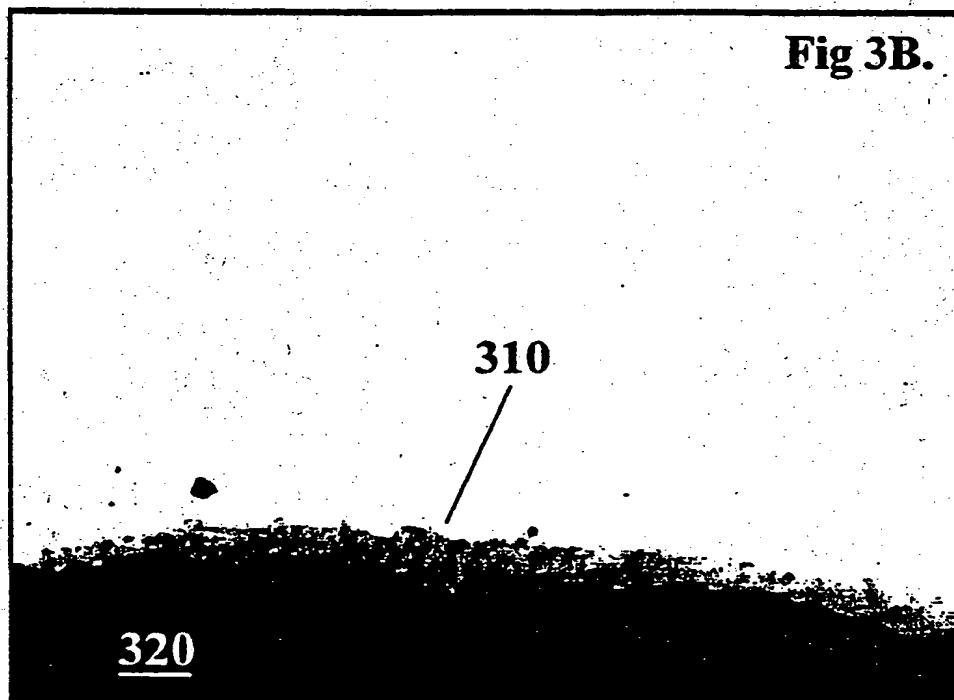


Fig 3B.



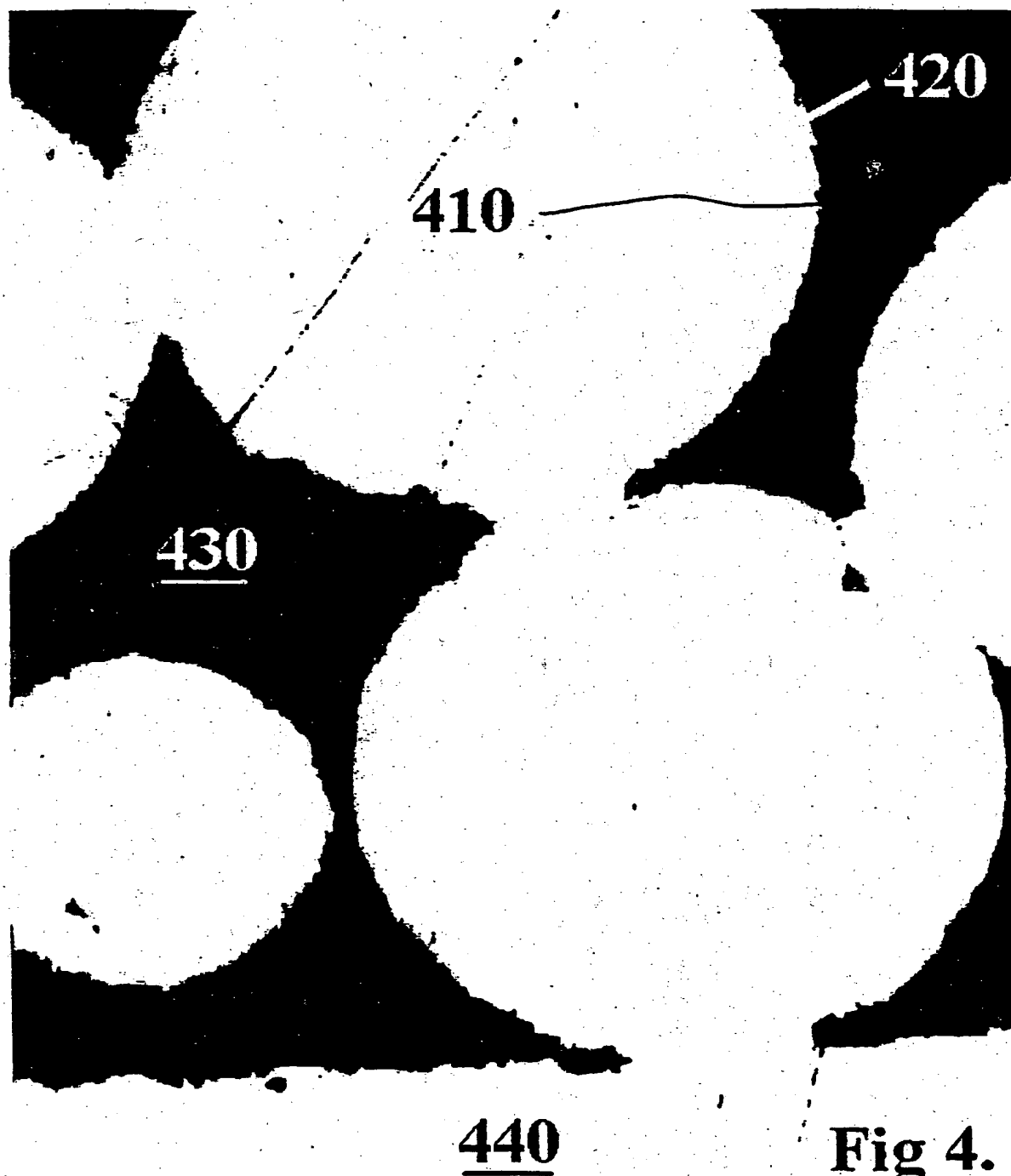


Fig 4.

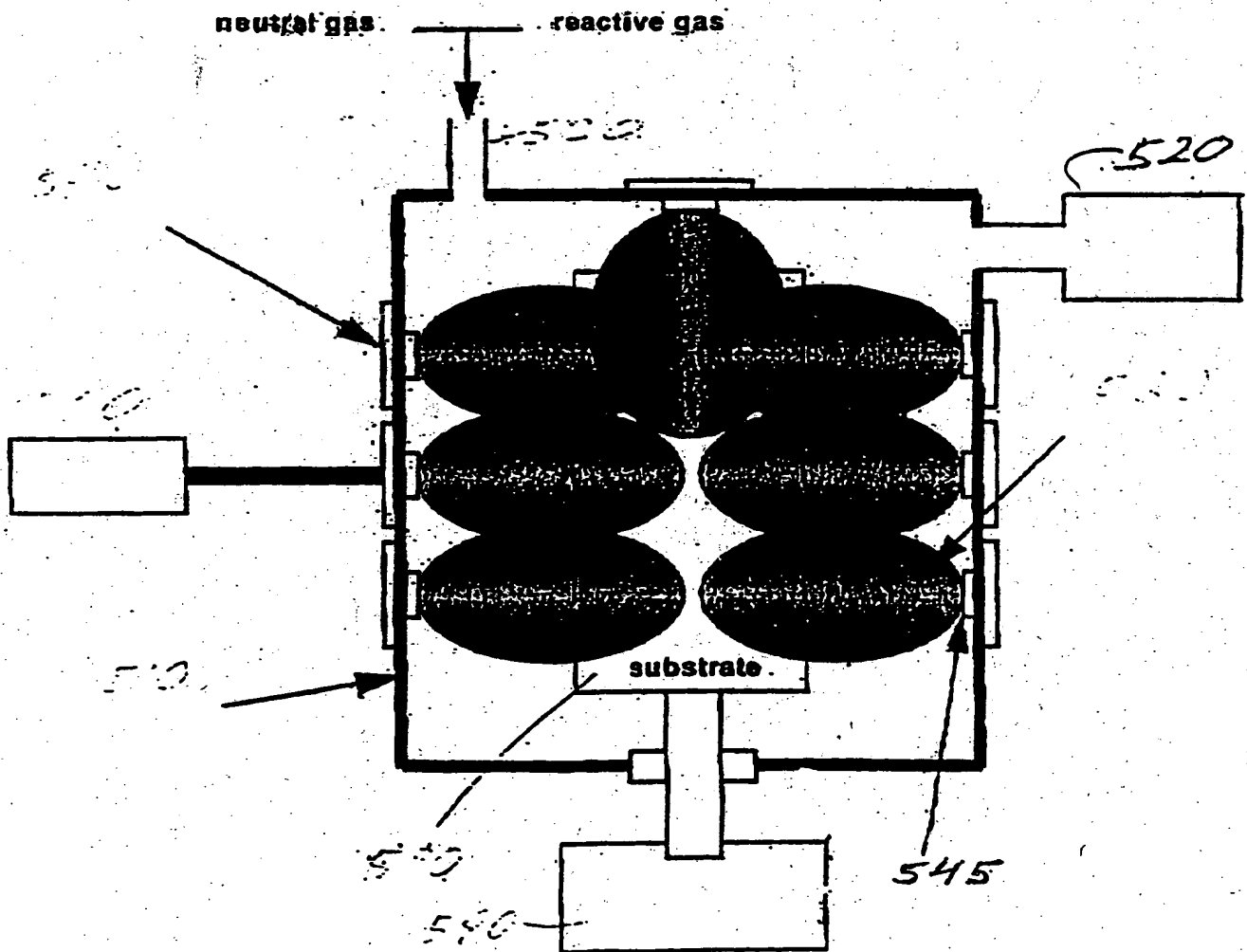
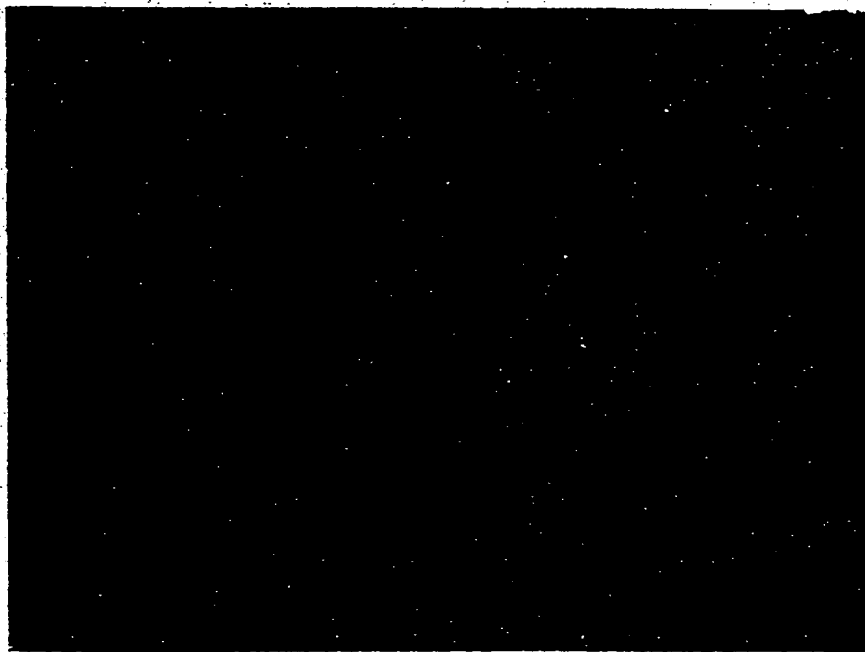


fig 5



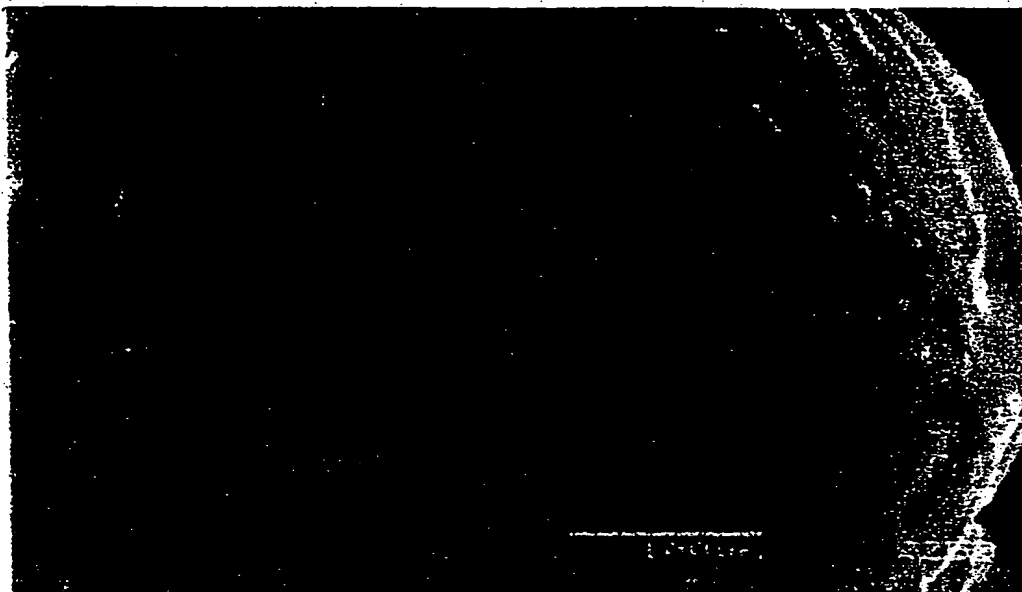
6A

SEM Micrograph of Group B at 300x after Electrochemical Test.

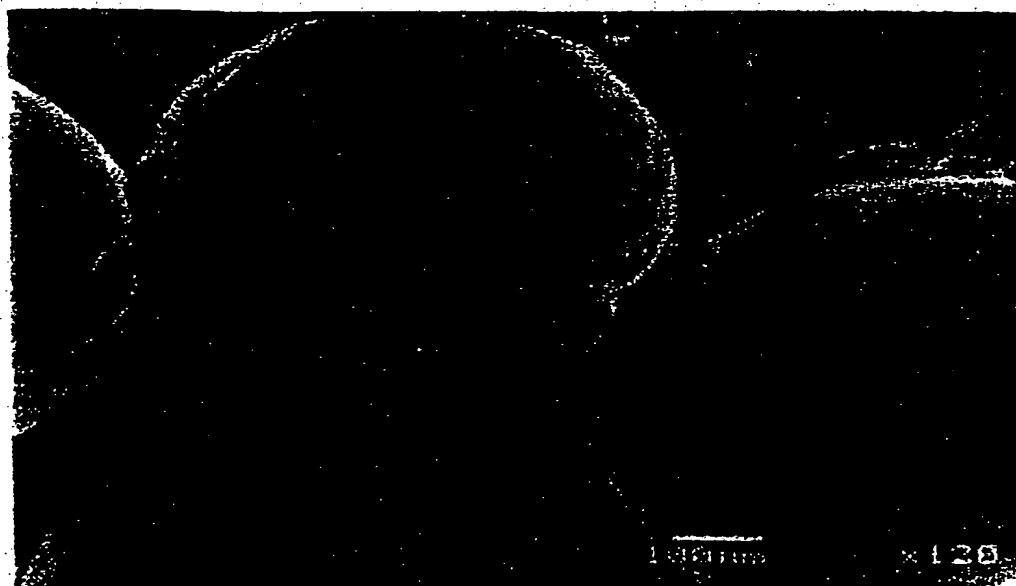


6B

SEM Micrograph of Group B at 700x after electrochemical test.



7A

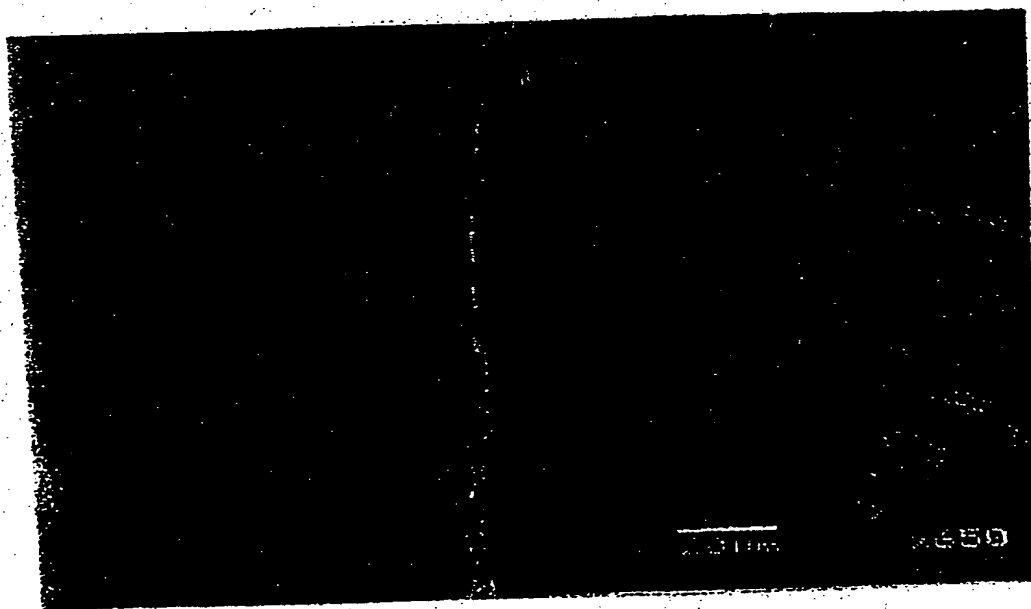


7B

SEM micrographs of the Hayes CrCoMo sample after ASTM F746 corrosion test indicates intergranular corrosion attacks.

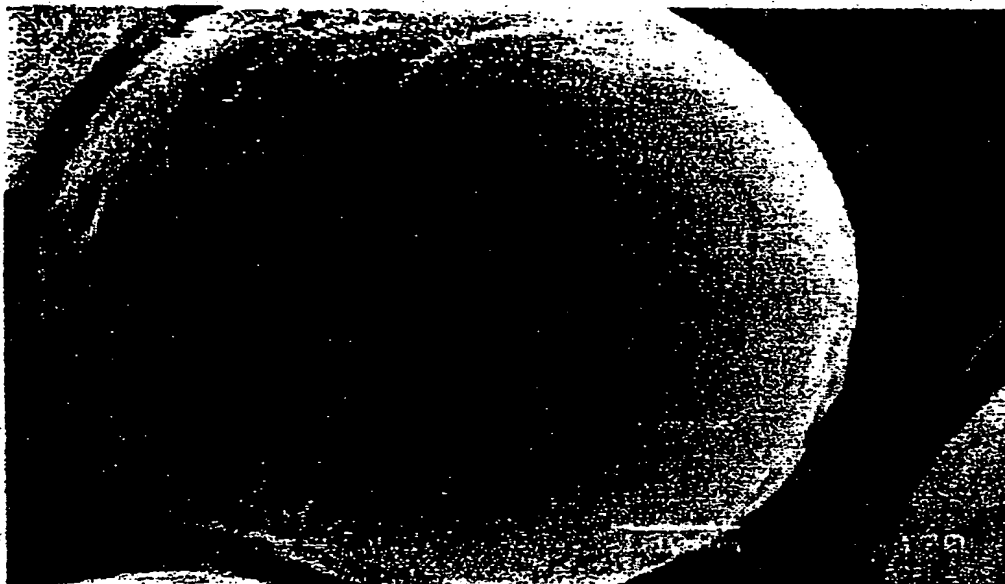


7C

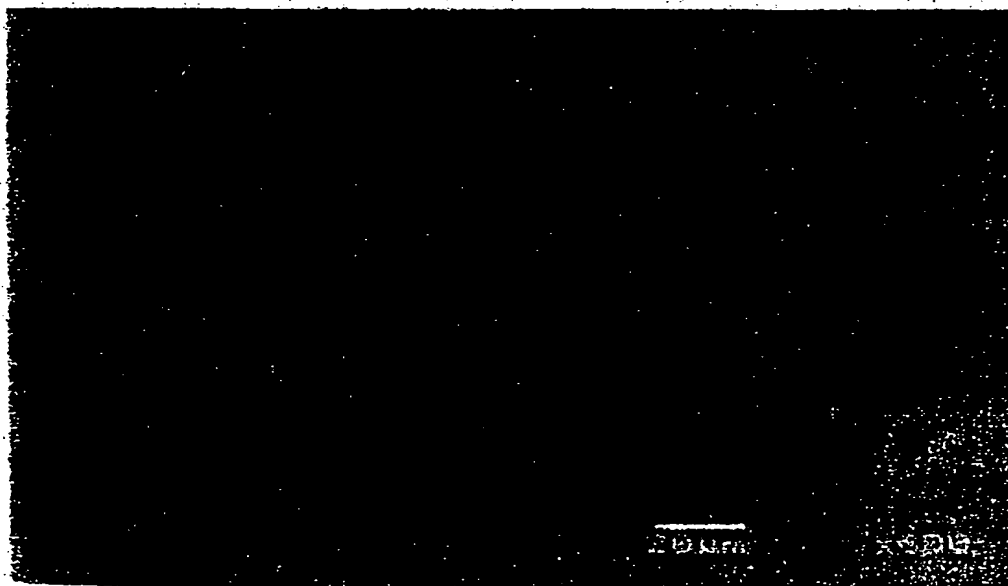


7D

SEM micrographs of the Hayes CrCoMo sample after ASTM F746 corrosion test exhibiting corrosion attacks.



8A



8B

SEM micrographs of the Hayes CrCo sample prior to any corrosion test.

INTERNATIONAL SEARCH REPORT

International Application No
PCT/US 99/08456

A. CLASSIFICATION OF SUBJECT MATTER
IPC 6 A61L27/00

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)
IPC 6 A61L C23C

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	<p>US 3 855 638 A (PILLIAR R) 24 December 1974 (1974-12-24)</p> <p>claims 1,4,10 column 8, line 27 - line 37</p> <p>— -/-</p>	<p>1-5, 8-13, 16-23, 26-31, 34-41, 44-49, 52-60, 63-68, 71-74</p>

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

*** Special categories of cited documents :**

- "A" document defining the general state of the art which is not considered to be of particular relevance
- "E" earlier document but published on or after the international filing date
- "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)
- "O" document referring to an oral disclosure, use, exhibition or other means
- "P" document published prior to the international filing date but later than the priority date claimed

- "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
- "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
- "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.
- "&" document member of the same patent family

Date of the actual completion of the international search

11 August 1999

Date of mailing of the international search report

19/08/1999

Name and mailing address of the ISA

European Patent Office, P.O. 5818 Patentlaan 2
NL - 2280 HV Rijswijk
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,
Fax: (+31-70) 340-3016

Authorized officer

Heck, G

INTERNATIONAL SEARCH REPORT

International Application No
PCT/US 99/08456

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 4 818 559 A (HAMA MASAOKI ET AL) 4 April 1989 (1989-04-04) column 2, line 58 - column 3, line 55 claims 1-3,5	1-6, 10, 12, 13, 16-24, 28, 30, 31, 34-42, 46, 48, 49, 52-55, 57-61, 65, 67, 68, 71-74
X	US 5 603 338 A (BEATY KEITH D) 18 February 1997 (1997-02-18) claims 1, 5, 6, 13	1-6, 10, 12, 13, 16-24, 28, 30, 31, 34-42, 46, 48, 49, 52-55, 57-61, 65, 67, 68, 71-74
X	WO 97 16137 A (BATCHELDER CLARENCE F ; SARKISIAN JAMES S (US)) 9 May 1997 (1997-05-09) page 4, line 13 - line 36 claims 16, 19	1-7, 10, 12, 16-25, 28, 30, 34-43, 46, 48, 52-55, 57-62, 65, 67, 71-74

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US 99/08456

Box I Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)

This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☒ Claims Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:
see FURTHER INFORMATION sheet PCT/ISA/210
2. ☐ Claims Nos.:
because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:
3. ☐ Claims Nos.:
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.
2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
3. ☐ As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:
4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:

Remark on Protest

- ☐ The additional search fees were accompanied by the applicant's protest.
- ☐ No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

International Application No. PCT/ US 99/08456

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box 1.1

Although claims 17, 18, 34, 36, 53, 54, 71, 72 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.

Continuation of Box 1.1

Rule 39.1(iv) PCT - Method for treatment of the human or animal body by surgery

INTERNATIONAL SEARCH REPORT

International Application No. PCT/US 99 08456

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

Continuation of Box I.1

Although claims 17, 18, 34, 36, 53, 54, 71, 72 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.

Continuation of Box I.1

Rule 39.1(iv) PCT - Method for treatment of the human or animal body by surgery

INTERNATIONAL SEARCH REPORT

information on patent family members

International Application No

PCT/US 99/08456

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
US 3855638 A	24-12-1974	CA 962806 A DE 2127843 A FR 2095854 A CH 540044 A GB 1316809 A	18-02-1975 16-12-1971 11-02-1972 28-09-1973 16-05-1973
US 4818559 A	04-04-1989	JP 8029150 B JP 62034558 A JP 62034559 A EP 0212929 A	27-03-1996 14-02-1987 14-02-1987 04-03-1987
US 5603338 A	18-02-1997	AU 4505196 A BR 9509934 A EP 0794745 A NO 972425 A WO 9616611 A US 5876453 A US 5863201 A	19-06-1996 27-01-1998 17-09-1997 28-05-1997 06-06-1996 02-03-1999 26-01-1999
WO 9716137 A	09-05-1997	AU 7456596 A	22-05-1997